

## EXHIBIT A

The following provides details for support for the amendments to Table 4 of the Specification and illustrates the amendments introduce no new matter.

In particular, the following provides sequence information as shown in originally filed Table 4 followed by the matching portion of preliminarily amended Table 4 (showing the proposed amendments).

### Specification as Filed for SEQ ID NOS: 49 and 54:

*Underlining in the original specification denotes support for amendments.*

Re- gion	Epi- tope	Clone	Lib <sup>a</sup>	Framework 1	CDR 3	Framework 4	
V <sub>H</sub>	1	C15	1	QVRLQSSGAELVIRPGASVKLSCKTSQYSPT	GIITFDYGGNYYAMN	WGQSTTVTASS	←SEQ ID NO: 49
		C9	1	-----	-----	-----	as originally filed
		1D5	2	E--VE-----N--A----	-----	-----	
		C1	1	-----	-----	-----	
		S25	1	-----	-----	-----	
		1H6	2	--Q-----V--I--A--T-I	-----	-----	
		1C9	2	--Q-E-----V--I--S--T-I	-----	-----	←SEQ ID NO: 54
		1E8	2	E-Q--E--PG--K-SQ-LS-T-TVT--I-	-----	-----	as originally filed
		1G7	2	E-Q--E--PG--K-SQ-LS-T-TVT--I-	-----	-----	

### Specification as Amended for SEQ ID NOS: 49 and 54:

V <sub>H</sub> Region		Sequence <sup>a</sup>			
Epitope 1		Framework 1	CDR 1	Framework 2	CDR 2
		Framework 3	CDR3	Framework 4	Seq ID No
C9	1	QVRLQSSGAELVIRPGASVKLSCKTSQYSPT MATLTVDKSSSFAYMQLASPTSEDSAVVYCAR	SYMMN GIYYVYGGNYYAMDY	WVKQGPQGLEMIG WGQSTTVTVSS	MIHPSNSEIRPNQKFN 49
<u>YY</u>					
1C9	2	QVRLQSSGAELVIRPGASVKLSCKTSQYSPT KATLTVDKSSSFAYMQLASPTSEDSAVVYCAR	DYAVH Rgkg	WVKQshaksLEWIG WGQSTTVTVSS	vistyygdadyNpkPkg 54

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### Specification as Filed for SEQ ID NO: 63:

Re- gion	Epi- tope	Clone	Lib*	Framework 1	CDR 1
	3	1A1	1	EVSLVESGGGLVQPGGSLRLSCAASGFTFS	DYVMS
		1E1	2	-----A-----	N-Q--
		C39	1	Q-Q-Q-----S-K-----L-----A-----	----
		C25	1	Q-Q-Q-----K-----L-----A-----	----Y
		2G5	2	-----K-----L-----A-----	S-A--
		3C3	2	-----K-----K-----L-----A-----	S-A--
		3F4	2	-----K-----L-----A-----	S-A--
		3H4	2	-----K-----PL-----A-----	S-A--

← SEQ ID NO: 63  
as originally filed

### Specification as Amended for SEQ ID NO: 63:

3F4	2	<u>E</u>	EVSLVESGGGLVQPGGSLRLSCAASGFTFS RFTTISADNAKINLYIQNSHLSEDTAMYYCAR	SYAMS slpydhv Dy	WVQGPshRLRWVA WQGTsVIVSS	TISDGGTfTYVCDaVKG 63
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### Specification as Filed for SEQ ID NOS: 68 and 69:

Re- gion	Epi- tope	Clone	Lib*	Framework 1	CDR 1
	3	1B3	2	EVQLQESGGGVQPGGSLRLSCAASGFTFS	SYAMH
		1C6	2	QI--LQ-----	----
		2B6	2	VKLVESGP-L-KPSQSLSLCTVTGYBIT-	D-AAM
		1G5	2	Q---Q--ABL---A-VNM--E---Y--T	---WTT ← SEQ ID NO: 68
		1H6	2	---Q--ABL_K--A-VNM--E---Y--T	---WTT ← SEQ ID NO: 69

as originally filed

### Specification as Amended for SEQ ID NOS: 68 and 69:

1G5	2		qVQLQqSGaeIvQPGaSVkmSCKASGyTft kaTltvDtsSasTaYmQlsSLtsEDsAVYYCAR	Wwtt <u>S</u> Elgd andy	WVkrPGqGLEWlg WQGTsVIVSS	dIypDsgstnynekfKa 68
1H6	2		<sup>K</sup> EVQLQqSGaeIvQPGaSVkmSCKASGyTft kaTltvDtsSasTaYmQlsSLtsEDsAVYYCAR	Wwtt <u>S</u> Elgd andy	WVkrPGqGLEWlg WQGTsVIVSS	dIypDsgstnynekfKa 69

# **EXHIBIT A**

## **Specification as Filed for SEQ ID NOS: 72-80:**

Clone	Lib	Framework 1	CDR 1	Framework 3	Framework 4	
C15	1	DIELTQSPAIMSASPGEKVIITC	SASS	SVPIRFGSGSGSTSYSLTISMEAEDESATYYC	FGGTKLELKR	SEQ ID NO: 72
C9	1	--D-----S-----I--	----	--A-----SV---A-----	--A-----I--	SEQ ID NO: 73
1D5	2	-----A-----I--	---S	--V-----S-----A-----	--G-----I--	SEQ ID NO: 74
C1	1	-----L-A-----I--	----	--V-----A-----	--A-----	SEQ ID NO: 75
S25	1	-----L-A-----I--	-V--E	--V-----S-----A-----	--A-----I--	SEQ ID NO: 76
1B6	2	-----SLAV-L-QRA-IS-	RA-ESV	--I-A-----R-DFT--INPV--D-V----	--A-----	SEQ ID NO: 77
1C9	2	-----SLAV-L-QRA-IS-	RA-ESV	--I-A-----R-DFT--RPV--D-V----	--S-----I--	SEQ ID NO: 78
1E8	2	-----	----	--A-----S-----A-----	--A-----	SEQ ID NO: 79
1G7	2	-----	----	--A-----S-----A-----	--A-----	SEQ ID NO: 80

as originally filed

## **Specification as Amended for SEQ ID NOS: 72-80:**

V <sup>h</sup> Region						
Epitope 1						
Clone	Lib	Framework 1 Framework 3	CDR 1 CDR 3	Framework 2 Framework 4	CDR 2	Seq ID NO
C15	1	DIELTQSPAIMSASPGEKVIITC G <sub>1</sub> SVPIRFGSGSGSTSYSLTISMEAEDESATYYC	SASS SVSNMY QQWSSYPFT	WYQKPGGSPRLLIY FGGTKLELKR	DTSNLAS	72
C9	1	DIELTQSPAIMSASPGEKVIITC G <sub>1</sub> SVPaRFGSGSGSTSYSLTISvMEAEDESATYYC	SASS SVSyMh QQyGgYPII	WfQQKPGtSPkpwIY FGAGTKLEIKR	DTSNLAS	73
1D5	2	DIELTQSPAIMSASPGEKVIITC G <sub>1</sub> SVPaRFGSGSGSTSYSLTISsMEAEDESATYYC	SASSs iSsSnIh QQWgSYPII	WYQKsGtSPkpwIY FGGTKLEIKR	DTSNLAS	74
C1	1	DIELTQSPAIMSASPGEKVIITC G <sub>1</sub> SVPaRFGSGSGSTSYSLTISrMEAEDESATYYC	SASS SVSyMY QQWSSYPIT	WYQKPGGSPRLLIY FGAGTKLELKR	DTSNLAS	75
S25	1	DIELTQSPAIMSASPGEKVIITC G <sub>1</sub> SVPaRFGSGSGSTSYSLTISsMEAEDESATYYC	SvSSs iSsSnIb QQWSSYPIT	WYQKsGtSPkpwIY FGAGTYLEIKR	DTSNLAS	76
1B6	2	DIELTQSPAslavSlQgraliSc G <sub>1</sub> SVPaRFGSGSGSrTdfLLTlnpVEADgVATYYC	ra <sup>1</sup> esvdsygnSfMh S QQanedPpT	WYQKPGGpPKLLIY FGAGTKLELKR	raSNLeS	77
1C9	2	DIELTQSPAslavSlQgraliSc G <sub>1</sub> SVPaRFGSGSGSrTdfLLTlnpVEADgVATYYC	ra <sup>1</sup> esvdsygnSfMh S QQanedPyT	WYQKPGGpPKLLIY FGG <sup>1</sup> TKLEIKR	raSNLeS	78
1E8	2	DIELTQSPAIMSASPGEKVIITC G <sub>1</sub> SVPaRFGSGSGSTSYSLTISsMEAEDESATYYC	SASS SVSyMh QQWSSnPII	WYQKsGtSPkpwIY FGAGTKLELKR	DTSKLAS	79
1G7	2	DIELTQSPAIMSASPGEKVIITC G <sub>1</sub> SVPaRFGSGSGSTSYSLTISsMEAEDESATYYC	SASS SVSyMh QQWSSnPII	WYQKsGtSPkpwIY FGAGTKLELKR	DTSKLAS	80

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### Specification as Filed for SEQ ID NOS: 81-88:

Clone	Lib <sup>o</sup>	Framework 1	CDR 1	CDR 3	Framework 4	
1A1	2	DIELTQSPASLAVSLGQRATISC	RASESVDSYGN <del>S</del> FMH	Q <del>QWSS</del> NPPT	FGSGTKLE <del>L</del> KR	SEQ ID NO: 81
1F1	2	DIELTQSPASLAVSLGQRATISC	RASESVDSYGN <del>S</del> FMH	Q <del>QWSS</del> NPPT	FGSGTKLE <del>L</del> KR	SEQ ID NO: 82
C39	1	DIELTQSPASLAVSLGQRATISC	RASESVDSYGN <del>S</del> FMH	Q <del>QWSS</del> NPPT	FGSGTKLE <del>L</del> KR	SEQ ID NO: 83
C25	1	DIELTQSPASLAVSLGQRATISC	RASESVDSYGN <del>S</del> FMH	Q <del>QWSS</del> NPPT	FGSGTKLE <del>L</del> KR	SEQ ID NO: 84
2G5	2	DIELTQSPASLAVSLGQRATISC	RASESVDSYGN <del>S</del> FMH	Q <del>QWSS</del> NPPT	FGSGTKLE <del>L</del> KR	SEQ ID NO: 85
3C3	2	DIELTQSPASLAVSLGQRATISC	RASESVDSYGN <del>S</del> FMH	Q <del>QWSS</del> NPPT	FGSGTKLE <del>L</del> KR	SEQ ID NO: 86
3F4	2	DIELTQSPASLAVSLGQRATISC	RASESVDSYGN <del>S</del> FMH	Q <del>QWSS</del> NPPT	FGSGTKLE <del>L</del> KR	SEQ ID NO: 87
3H4	2	DIELTQSPASLAVSLGQRATISC	RASESVDSYGN <del>S</del> FMH	Q <del>QWSS</del> NPPT	FGSGTKLE <del>L</del> KR	SEQ ID NO: 88

as originally filed

### Specification as Amended for SEQ ID NOS: 81-88:

1A1	2	DIELTQSPASLAVSLGQRATISC	RASESVDSYGN <sup>H</sup> SFMH	WYQQKPGQPPKLLIY	LASNLES	
		GVPARFSGSGSKTDFTLTIDPEADDAATYYC	Q <del>QWSS</del> NPPT NNEDPYT	FGSGTKLE <del>L</del> KR	81	
1F1	2	DIELTQSPASLAVSLGQRATISC	RASESVDSYGN <sup>G</sup> SFMH	WYQQKPGQPPKLLIY	LASNLES	
		GVPARFSGSGSKTDFTLTIDPEADDAATYYC	Q <del>QWSS</del> NPPT NNEDPY	FGSGTKLE <del>L</del> KR	82	
C39	1	DIELTQSPASLAVSLGQRATISC	RASESVDSYGN <sup>I</sup> SFMH	WYQQKPGQPPKLLIY	LASNLES	
		GVPARFSGSGSKTDFTLTIDPEADDAATYYC	Q <del>QWSS</del> NPPT NNEDPY	FGSGTKLE <del>L</del> KR	83	
C25	1	DIELTQSPASLAVSLGQRATISC	RASESVDSYGN <sup>G</sup> SFMg	WYQQKPGQPPKLLIY	rASNLEp	
		GVPARFSGSGSKTDFTLTIDPEADDAATYYC	Q <del>QWSS</del> NPPT NNEDPY	FGSGTKLE <del>L</del> KR	84	
2G5	2	DIELTQSPAlmsaSpGekvtctC	RASs svSyMG	WYQQKPGGlsPklwIY	stSNLaS	
		GVPARFSGSGSgTsysLTlarmEAeDAATYYC	Q <del>QWSS</del> NPPT	FGSGGqagnKS	85	
3C3	2	DIELTQSPAlmsaSpGekvtctC	RASESVDSYGN <sup>I</sup> SFMg	WYQQKPGGlsPklwIY	stSNLaS	
		GVPARFSGSGSgTsysLTlarmEAeDAATYYC	Q <del>QWSS</del> NPPT	FGSGGqagnKS	86	
3F4	2	DIELTQSPAlmsaSpGekvtctC	RASs svSyMy	WYQQKPGGlsPklwIY	stSNLaS	
		GVPARFSGSGSgTsysLTlarmEAeDAATYYC	Q <del>QWSS</del> NPPT YPPT	FGSGTKLE <del>L</del> KR	87	
3H4	2	DIELTQSPAlmsaSpGekvtctC	RASs vssSyIG	WYQQKPGGlsPklwIY	stSNLaS	
		GVPARFSGSGSgTsysLTlarmEAeDAATYYC	Q <del>QWSS</del> NPPT	FGSGTKLE <del>L</del> KR	88	

YPP I

## EXHIBIT A

### Specification as Filed for SEQ ID NOS: 94 and 95:

Clone	Sequence <sup>a</sup>	
	CDR 2	Framework 3
1F3	SASRLAS	GVPSRFSGSGSGTSYSLTISSVEARDAATYYC
2E8	ET-----	---A-----GAS---V-----

← SEQ ID NO: 94  
 ← SEQ ID NO: 95  
 as originally filed

### Specification as Amended for SEQ ID NOS: 94 and 95:

SL					
1F3	2	DIELTQSPASMSASPGEKVTMT	KATSS	VSSSYLH	WYQQKSGASPKLWIY
		GVPSRFSGSGSGTSYSLTISSVEARDAATYYC	QYIGYFTT	FGGSKLEIKR	SASRLAS
					94
2E8	2	DIELTQSPtMaASPGEKItiTC	sAsSS	igSnYLN	WYQQKpGfSPKLIIY
		GVPSRFSGSGSGTSYSLTIgaSAEDvATYYC	QQgsaIFTT	FGGSKLEIKR	★tSNLAS
					<u>R</u> 95

M

## EXHIBIT B

The following illustrates that the amendments to the claims to correct the sequence identifiers do not introduce new matter.

The relevant parts of Table 11 on page 85-88, paragraph [0241] of the Specification as originally filed are reproduced below.

[0241] Table 11 amino acid sequences for affinity matured and/or modified antibodies.

Heavy Chains				
Clone	Framework 1	CDR1	Framework 2	CDR2
huC25	QVQLQESGGGLVQPGGSLRLSC AASGFTFS (SEQ ID	DYYMY (SEQ ID NO: 87)	WVRQAPGKGLEW VA (SEQ ID	TTSDGGSYTYYPD SVRG (SEQ ID

	NO: 86)		NO: 88)	NO: 89)
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...

Heavy Chains cont'd				
	Framework 3	CDR3	Framework 4	
huC25	RFTISRDNKNTLYLQMNLSLRA EDTAMYYCSR (SEQ ID NO: 126)	YRYDDAMDY (S EQ ID NO: 127)	WGQGTLLVTVSS (SEQ ID NO: 128)	

...

Light Chains				
Clone	Framework 1	CDR1	Framework 2	CDR2
huC25	EIVLTQSPATLSLSPGERATIS	RASESVDSYGH	WYQQKPGQAPRL	RASNLEP (SEQ
	C (SEQ ID NO: 156)	SPMQ (SEQ ID NO: 157)	LIY (SEQ ID NO: 158)	ID NO: 159)

...

Light Chains cont'd.				
Clone	Framework 3	CDR3	Framework 4	
huC25	GIPARFSGSGSGTDFTLTISL EPEDFAVYYC (SEQ ID NO: 196)	QQSNEDPFT (S EQ ID NO: 197)	FGQGTKVEIKR (SEQ ID NO: 198)	

## EXHIBIT B

The relevant parts of Table 11 as amended pursuant to the **Preliminary Amendment** filed April 19, 2004 are reproduced below.

At pages 85-88, amend Table 11 as follows:

[0241] Table 11 amino acid sequences for affinity matured and/or modified antibodies.

Heavy Chains				
Clone	Framework 1	CDR1	Framework 2	CDR2
huC25	QVQLQESGGGLVQPGGSLRLSC AASGFTFS (SEQ ID NO: <del>86</del> 138)	DYYMY (SEQ ID NO: <del>87</del> 139)	WVRQAPGKGLEW VA (SEQ ID NO: <del>88</del> 140)	TISDGGSYTYYPD SVKG (SEQ ID NO: <del>89</del> 141)

...

Heavy Chains cont'd				
	Framework 3	CDR3	Framework 4	
huC25	RFTISRDN SKNTLYLQMNSLRA EDTAMY YCSR (SEQ ID NO: <del>126</del> 178)	YRYDDAMDY (S EQ ID NO: <del>127</del> 179)	WGQGT LVT VSS (SEQ ID NO: <del>128</del> 180)	

...

Light Chains				
Clone	Framework 1	CDR1	Framework 2	CDR2
huC25	EIVLTQSPATLSLSPGERATIS C (SEQ ID NO: <del>156</del> 208)	RASESVDSYGH SFMQ (SEQ ID NO: <del>157</del> 209)	WYQQKPGQAPRL LIY (SEQ ID NO: <del>158</del> 210)	RASNLEP (SEQ ID NO: <del>159</del> 211)

...

Light Chains cont'd.				
Clone	Framework 3	CDR3	Framework 4	
huC25	GIPARFSGSGSGTDFTLTISL EPEDFAVYYC (SEQ ID NO: <del>196</del> 248)	QQSNEDPFT (SEQ ID NO: <del>197</del> 249)	FGQGTRVEIKR (SEQ ID NO: <del>198</del> 250)	

## EXHIBIT B

The sequence information of SEQ ID NOS: 139, 141, 179, 209, 211, and 249, as shown in the enclosed Sequence Listing, as well as their one letter code sequences are as follows.

### As shown in the Sequence Listing:

### One-Letter Code:

### Sequence in the Specification as Amended in the Preliminary Amendment:

<210> 139  
 <211> 5  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> single chain antibody fragment

<400> 139  
 Asp Tyr Tyr Met Tyr  
 1 5

SEQ ID NO: 139  
 DYYMY

DYYMY (SEQ  
 ID  
 NO: ~~87~~139)

<210> 141  
 <211> 17  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> single chain antibody fragment

<400> 141  
 Thr Ile Ser Asp Gly Gly Ser Tyr Thr Tyr Tyr  
 Pro Asp Ser Val Lys  
 1 5 10 15  
 Gly

SEQ ID NO: 141  
 TISDGGSYTYYPDSVKG

TISDGGSYTYYPD  
 SVKG (SEQ ID  
 NO: ~~89~~141)

<210> 179  
 <211> 9  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> single chain antibody fragment

<400> 179  
 Tyr Arg Tyr Asp Asp Ala Met Asp Tyr  
 1 5

SEQ ID NO: 179  
 YRYDDAMDY

YRYDDAMDY (S  
 EQ ID  
 NO: ~~127~~179)



**EXHIBIT B**

<210> 209  
<211> 15  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> single chain antibody fragment

<400> 209  
Arg Ala Ser Glu Ser Val Asp Ser Tyr Gly His  
Ser Phe Met Gln  
1 5 10 15

SEQ ID NO: 209  
RASESVDSYGH~~S~~FMQ

RASESVDSYGH  
SFMQ(SEQ ID  
NO: ~~157~~209)

<210> 211  
<211> 7  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> single chain antibody fragment

<400> 211  
Arg Ala Ser Asn Leu Glu Pro  
1 5

SEQ ID NO: 211  
RASNLEP

RASNLEP(SEQ  
ID NO: ~~159~~211)

<210> 249  
<211> 9  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> single chain antibody fragment

<400> 249  
Gln Gln Ser Asn Glu Asp Pro Phe Thr  
1 5

SEQ ID NO: 249  
QQSNEDPFT

QQSNEDPFT  
(SEQ ID  
NO: ~~197~~249)